1) Publication number:

0 370 656 A1

	_	
•	-	١
(12	

EUROPEAN PATENT APPLICATION

- (1) Application number: 89311507.1
- (a) Int. Ci.5: A61K 37/02, //(A61K37/02, 31:52)

- 2 Date of filing: 07.11.89
- (3) Priority: 09.11.88 US 269276
- ② Date of publication of application: 30.05.90 Bulletin 90/22
- Designated Contracting States:
 ES GR

- 7) Applicant: SCHERING CORPORATION 2000 Gailoping Hill Road Kenilworth New Jersey 07033(US)
- inventor: Bonnem, Eric M. 622 Belvidere Avenue Plainfield New Jersey 07062(US)
- Representative: Ritter, Stephen David et al Mathys & Squire 10 Fleet Street London EC4Y 1AY(GB)
- Treatment of myelosuppression associated with acquired immune deficiency.
- Pharmaceutical compositions and methods of treatment are useful for treating myelosupression and viral infections associated with AIDS, particularly cytomegalovirus retinitis. Such compositions and treatments utilize an antiviral effective amount of 9-(1,3-dihydroxy-2-propoxymethyl) guanine (DHPG) in combination with an anti-myelosuppressive effective amount of granulocyte macrophage colony stimulating factor (GM-CSF). The use of DHPG and of GM-CSF for the preparation of a pharmaceutical composition for use in this combination treatment as well as kits containing pharmaceutical compositions containing DHPG and GM-CSF are also disclosed.

EP 0 370 656 A1

EP0370656

- 1. A pharmaceutical composition comprising an antiviral effective amount of DHPG in combination with an anti-myelosuppressive effective amount of GM-CSF.
- 2. The pharmaceutical composition as claimed in claim 1 wherein the antiviral effective amount of DHPG is an amount sufficient to lessen the symptoms associated with cytomegalovirus retinitis.
- 3. The pharmaceutical composition as claimed in claims 1 or 2 wherein the antimyelosuppressive effective amount of GM-CSF is an amount sufficient to raise the white blood count to at least about 2000 per cm.
- 4. The pharmaceutical composition as claimed in anyone of claims 1 to 3 wherein the antimyelosuppressive effective amount of GM-CSF is an amount sufficient to raise the white blood cell count to from about 2000 to about 10,000 per cm.
- 5. The pharmaceutical composition as claimed in anyone of claims 1 to 4 wherein the antimyelosuppressive effective amount of GM-CSF is from about 3 to about 15 mu g per Kg per day.
- 6. The pharmaceutical composition as claimed in anyone of claims 1 to 5 further comprising an appropriate solution for intraveneous infusion.
- 7. A method of treating a patient suffering from myelosuppression and a viral infection associated with AIDS comprising administering to such a patient an antiviral effective amount of DHPG and an anti-myelosuppression effective amount of GM-CSF.
- 8. A method of treating a patient as claimed in claim 7 wherein a pharmaceutical composition of anyone of claims 1 to 6 is administered to the patient.
- 9. The method of claim 7 wherein administration of the DHPG is intravenous and GM-CSF is subcutaneous.
- 10. The method of claim 7 wherein administration of the DHPG is intravenous and GM-CSF is intraveneous.
- 11. A method of treating a patient suffering from myelosuppression and cytomegalovirus retinitis associated with AIDS comprising administering to such a patient an antiviral effective amount of DHPG and an anti-suppressive effective amount of GM-CSF.
- 12. The use of DHPG for the preparation of a pharmaceutical composition for use in a combined therapy for treating a patient suffering from myelosuppression and a viral infection by administering DHPG in association with GM-CSF.
- 13. The use of GM-CSF for the preparation of a pharmaceutical composition for use in a

combined therapy for treating a patient suffering from myelosuppression and a viral infection by administering GM-CSF in association with DHPG.

- 14. The use of DHPG in association with GM-CSF for the preparation of a pharmaceutical composition for treating a patient suffering from myelosuppression in association with a viral infection, said pharmaceutical composition comprising DHPG and GM-CSF together with pharmaceutically acceptable carriers therefor.
- 15. The use of DHPG in combination GM-CSF for treating a patient suffering from myelosuppression and a viral infection.
- 16. A process for the preparation of a pharmaceutical composition as claimed in anyone of claims 1 to 6 with comprises admixing DHPG and GM-CSF with pharmaceutically acceptable carriers therefor.
- 17. A kit for use in treating patients suffering from myelosuppression and a viral infection, said kit comprising a pharmaceutical package having a first container and a second container, wherein the first container comprises a pharmaceutical composition containing an antiviral effective amount of DHPG and the second container comprises a pharmaceutical composition containing anti-myelosuppressive amount of GM-CSF.